

Sample Application 2

Cytochrome P450 Inhibition

The Cytochrome P450 inhibition was performed according to the publication of Crespi et al. (Anal. Biochem., 248, 188-190 (1997)) with use of the baculovirus/insect cell-expressed, human Cytochrome P 450 isoenzyme (3A4).

The results are presented in the following table.

| Example No. | VEGFR II (KDR) | Cytochrome P450 |
|-------------|----------------|-----------------|
| | [nM] | Isoenzyme 3A4 |
| | | <u>μΜ</u> |
| 2.54 from | 5 | 3.6 |
| WO 00/27819 | | |
| 38 from | 180 | 4.6 |
| WO 00/27820 | | |
| 1.14 | 52 | >30 |
| 3.24 | 12 | 14 |
| 3.30 | 10 | 5.5 |
| 6.2 | 41 | >30 |
| 6.22 | 24 | 10 |
| 6.27 | 8 | 10 |
| 6.32 | 65 | . 11 |

The superior action of the compounds according to the invention compared to the known compounds can be seen clearly from the result.